The Swedi h Patent Office

PCT Int mational Application

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- 1. Ljungan picornavirus, comprising in the non-coading region of its viral genome a nucleotide sequence corresponding to a cDNA sequence selected from the group consisting of
- SEQ ID NO: 1 (Ljungan 87-012) 10

AGTCTAGTCT TATCTTGTAT GTGTCCTGCA CTGAACTTGT TTCTGTCTCT 50 GGAGTGCTCT ACACTTCAGT AGGGGCTGTA CCCGGGCGGT CCCACTCTTC 100 ACAGGAATCT GCACAGGTGG CTTTCACCTC TGGACAGTGC ATTCCACACC 150 CGCTCCACGG TAGAAGATGA TGTGTGTCTT TGCTTGTGAA AAGCTTGTGA 200 AAATCGTGTG TAGGCGTAGC GGCTACTTGA GTGCCAGCGG ATTACCCCTA 250 GTGGTAACAC TAGC

and homologous sequences having at least 75% homology to the SEQ ID NO: 1, and further, causing mammalian disease.

- 2. Ljungan picornavirus according to claim 1, wherein said homologous sequences have at least 80%, at least 85% or at least 90% homology to the SEQ ID NO: 1.
  - 3. Ljungan picornavirus according to claim 2, wherein said homologous sequence is one of
- SEQ ID NO:2 (Ljungan 174F) 30

AGTCTAGTTT CATTCTGTGT GTGTTTGGCA CTGAAATTAT TTCTGTCTCT 50 GGGGTGCTTT ACACTTCAGT AGGGGCTGTA CCCGGGCGGT CCCACTCTTC 100 ACAGGAATNT GCACAGGTGG CTTTCACCTC TGGACAGTGC ATTCCACACC 150 CGCTCCACAG TAGAAGATGA TGTGTGTCTT TGCTTGTGAA AAGCTTGTGA 200 35 AAATCGTGTG TAGGCGTAGC GGNTACTTGA GTGCCAGCGG ACNACCCCTA 250 **GTGGTAACAC TAGC** 

and

SEQ ID NO:3 (Ljungan 145SL). 40

AGTTTGGTTC TCTCTTGAGT GTGTTTTGTG TTAGCATAAT TTCTGTCTCT 50
AGAGTGCTTT ACACTCTAGT AGGGGCTGTA CCCGGGCGGT CCCACTCTTC 100
ACAGGAATCT GCACAGGTGG CTTTCACCTC TGGACAGTGC ATTCCATACC 150
CGCTCCACAA TAGAAGATGA TGTATATCTT TGTTTGTGAA ATGCTCATGA 200
AACGTGTGTG TAGGCGTAGC GGCTACTTGA ATGCCAGCGG AACCCCCCTA 250
GTGGTAACAC TAGC.

4. Protein comprising an amino acid sequence selected from the group consisting of

SEQ ID NO: 4 ( partial structural protein of Ljungan 145SL)

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Lys Asp Leu Met Glu Ile Ala Arg Met Pro Ser Val Tyr Lys Gly Glu 10 Arg Thr Glu Pro Gly Gly Thm Asn Gly Tyr Phe Gln Trp Ser His Thr 20 25 30 His Ser Pro Ile Asn Trp V∉l Phe Asp Gly Gly Ile His Leu Glu Asp 35 Met Pro Asn Leu Asn Leu Phe Ser Ser Cys Tyr Asn Tyr Trp Arg Gly 55 Ser Thr Val Leu Lys Leu Thr Val Tyr Ala Ser Thr Phe Asn Lys Gly 80 65 75 Arg Leu Arg Met Ala Phe Phe Pro Ile Met Met Gln Gly Thr Gln Arg 85 90 Leu Phe Met Val Cys Asp Ile Gly Leu Asn Asn Lys Lys His Lys Cys 105 Thr Phe Glu Met Thr Ile Pro Tyr Thr Trp Gly Asn Trp Met Arg Pro 115 120 125 Thr Arg Gly Ser Vall Ile Gly Trp Leu Arg Ile Asp Val Leu Asn Arg 130 135 140 Leu Thr Tyr Asn \$er Ser Pro Asn Ala Val Asn Cys Ile Leu Gln 150 155

Val Lys Met Gly Asn Asp Ala Lys Phe Met Val Pro Thr Thr Ser Asn
165 170 175

Ille Val Trp ,

and homologous sequences having at least 75% homology to the SEQ ID NO: 4,

and antigenid fragments of the sequences.

- 5. Antiserum or antibody directed against a structural protein of the virus according to any one of claims 1-3.
- 6. Antigen comprising at least a part of a structural protein of the picornavirus according to any one of claims 1-3.

7. Diagnostic kit comprising at least one member from the group consisting of an antiserum or antibody according to claim 5 or an antigen-binding part thereof, an antigen according to claim 6 or an antibody-binding part thereof, one or several probes designed with respect to the genome of the virus according to any one of claims 1-3,

and

one or several primers designed with respect to the genome of the virus according to any one of claims 1-3.

8. Vaccine having as an immunizing or neutralizing component a member selected from the group consisting of

a) the virus according to any one of claims 1-3,

- b) the virus according to any one of claims 1-3 in attenuated form,
- c) the virus according to any one of claims 1-3 in killed form,
- d) an antigen according to claim 6, including a subunit of the virus according to any one of claims 1-3, and
- e) DNA corresponding to the genomic RNA of the virus according to any one of claims 1-3.

9. Vaccine according to claim 8 which additionally comprises an adjuvant.

10. Ljungan picornavirus according to any one of the claims 1-3, optionally in attenuated or killed form, an antiserum or antibody according to claim-5 or an antigen according to claim-6 for use in a medicament.

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11. Use of a Ljungan picornavirus according to any one of the claims 1-3, optionally in attenuated or killed form, an antiserum or antibody according to claim 5 or an antigen according to claim 6, in the preparation of a medicament for prophylactic or therapeutic treatment of a disease caused by said virus.

12. Use according to claim 11, wherein the disease caused by said virus is one of Myocarditis, Cardiomyopathia, Guillain Barré Syndrome, and Diabetes Mellitus, Multiple Sclerosis, Chronic Fatigue Syndrome, Myasthenia Gravis, Amyothrophic Lateral Sclerosis, Dermatomyositis, Polymyositis, Spontaneous Abortion, and Sudden Infant Death Syndrome.

- 13. Method of prophylactic or therapeutic treatment of a disease caused by a virus according to any one of the claims 1-3 in a mammal, including human, which comprises administering to said mammal a prophylactically or therapeutically effective amount of a medicament comprising as an active ingredient a member of the group consisting of
- a) the virus according to any one of claims 1-3,
- b) the virus according to any one of claims 1-3 in attenuated form,
- c) the virus according to any one of claims 1-3 in killed form,
- d) an antigen according to claim 6, including a subunit of the virus according to any one of claims 1-3, and
- e) DNA corresponding to the genomic RNA of the virus according to any one of claims 1-3.

14. Method according to claim 13, wherein the disease caused by said virus is one of Myocarditis, Cardiomyopathia, Guillain Barré Syndrome, and Diabetes Mellitus, Multiple Sclerosis, Chronic Fatigue Syndrome, Myasthenia Gravis, Amyothrophic Lateral Sclerosis, Dermatomyositis, Polymyositis, Spontaneous Abortion, and Sudden Infant Death Syndrome.

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AMENDED SHEET